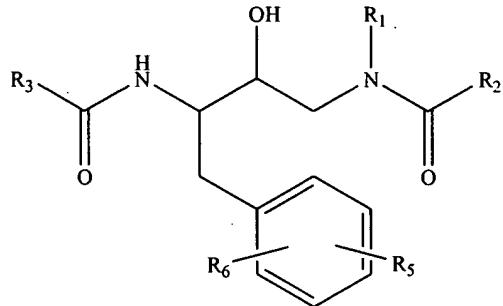


Amendments to the Claims:

This listing of claims will replace all prior versions, and listings of claims in the application:

Listing of Claims:

1 1 (currently amended): A method for modulating the processing of an amyloid
2 precursor protein (APP), said method comprising contacting a composition containing said APP
3 with an aspartyl protease inhibitor having the general formula:
4



5 wherein:

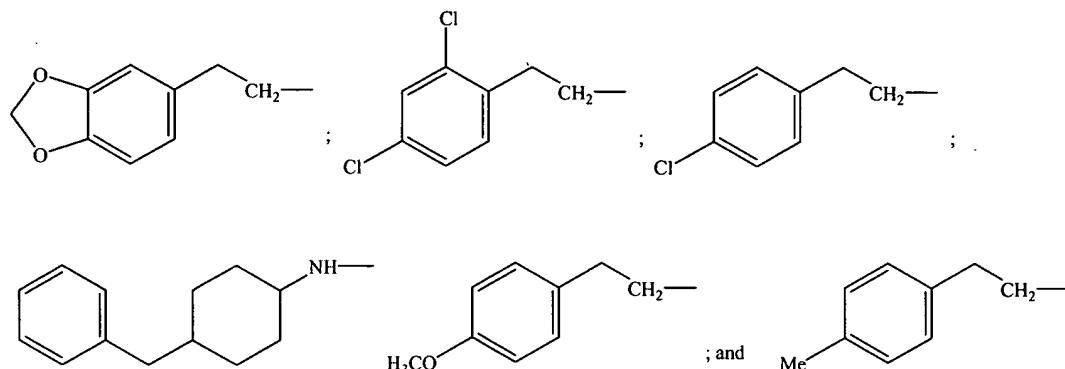
6 R₁, R₂ and R₃ are members independently selected from the group consisting of
7 alkyl, substituted alkyl, aryl, substituted aryl, arylalkyl, substituted
8 arylalkyl, aryloxyalkyl, substituted aryloxyalkyl, heteroaryl, substituted
9 heteroaryl, heteroarylalkyl, substituted heteroarylalkyl, heterocycles,
10 substituted heterocycles, heterocyclicalkyl and substituted
11 heterocyclicalkyl; and

12 R₅ and R₆ are independently selected from the group consisting of hydrogen,
13 halogen, alkyl, substituted alkyl, aryl, substituted aryl, arylalkyl,
14 substituted arylalkyl, aryloxyalkyl and substituted aryloxyalkyl; or R₅ and
15 R₆ and the carbons to which they are bound join to form an optionally

17 substituted carbocyclic or heterocyclic fused ring system having a total of
18 9- or 10-ring atoms within said fused ring system.

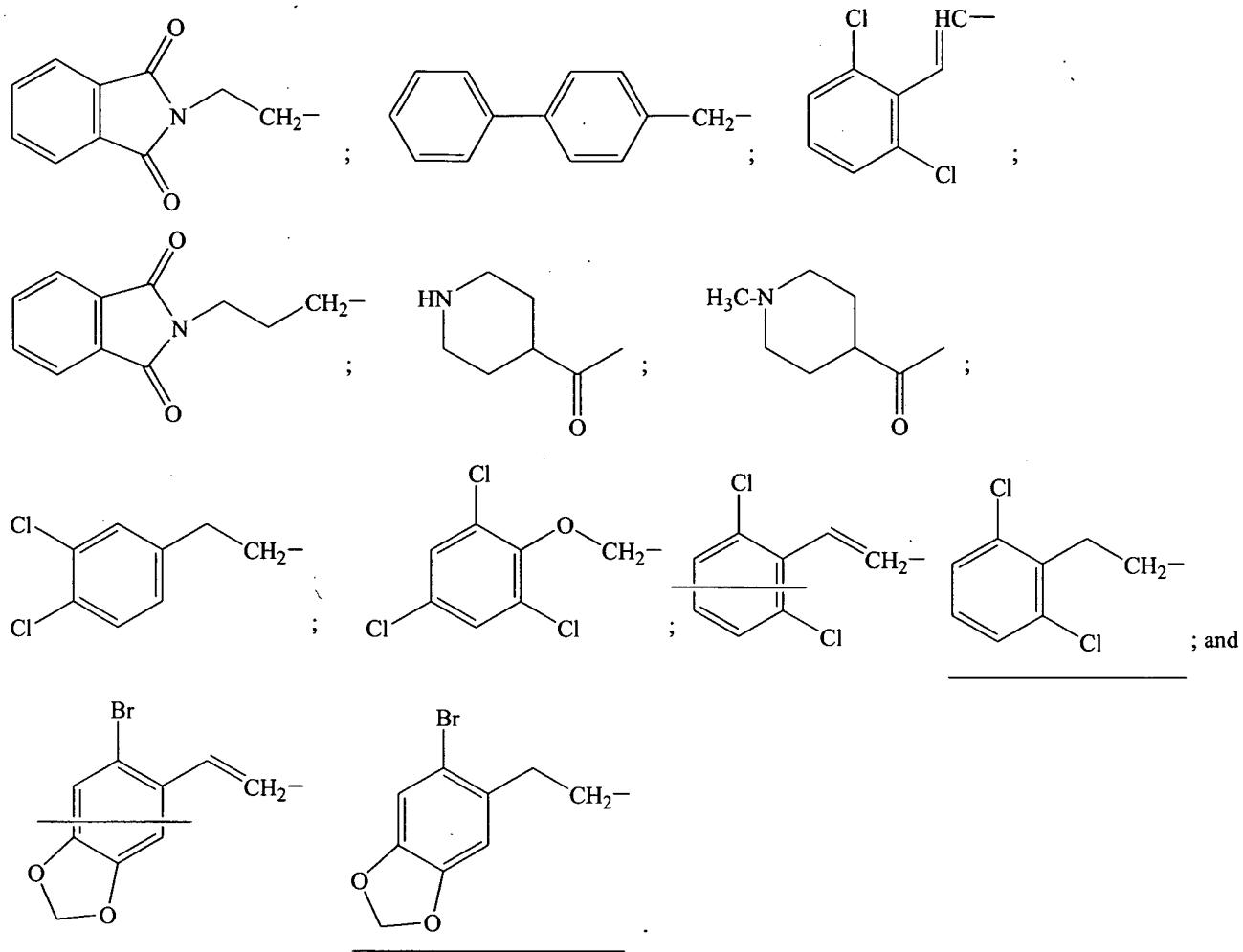
2 (original): The method according to claim 1, wherein:
R₁ is a member selected from the group consisting of substituted alkylaryl, substituted aryl, substituted alkyl and substituted heterocyclic groups.

3 (original): The method according to claim 2, wherein:
R₁ is a member selected from the group consisting of:



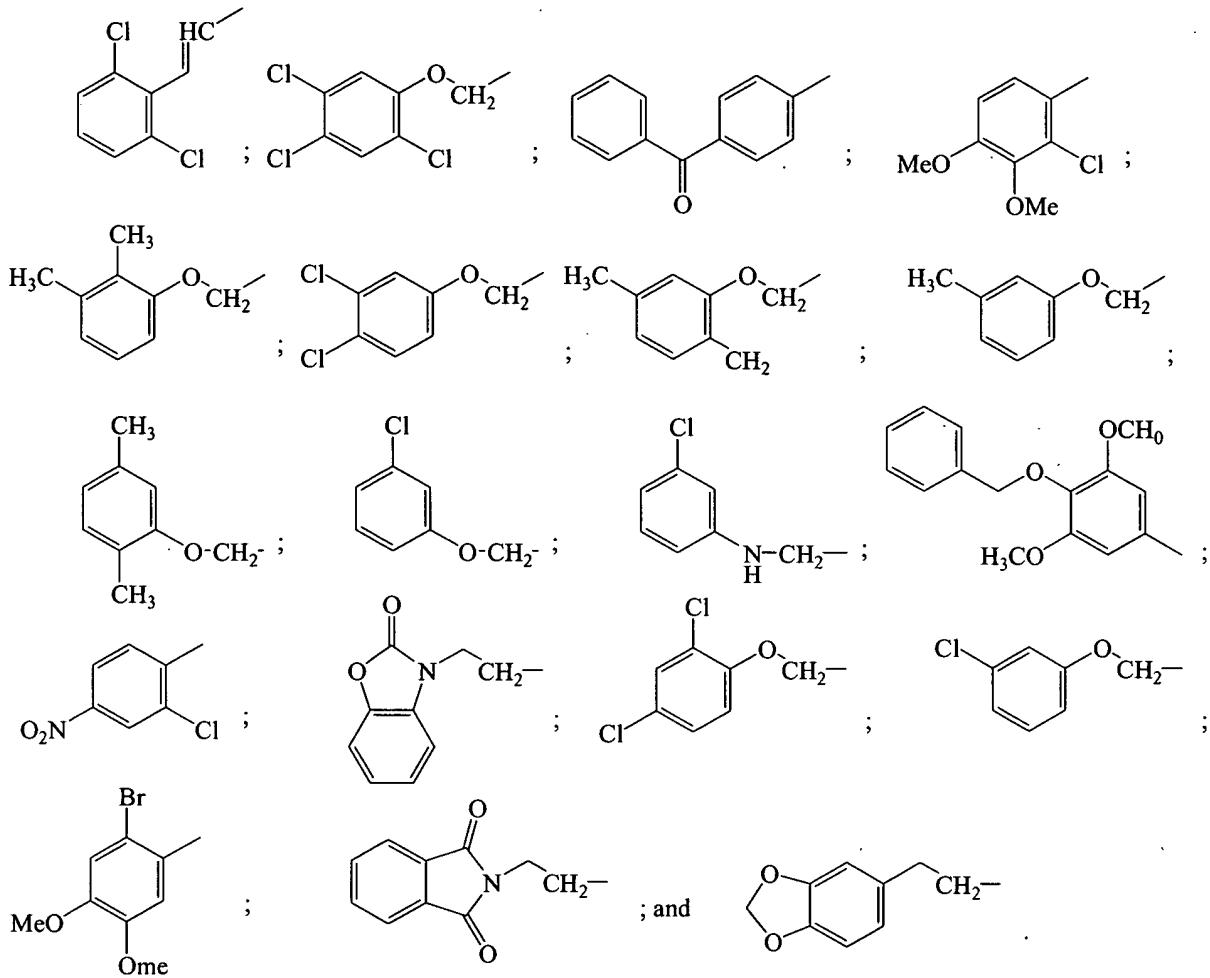
4 (original): The method according to claim 1, wherein:
R2 is a member selected from the group consisting of substituted alkyl,
and substituted heterocyclic groups.

5 (currently amended): The method according to claim 4, wherein R₂ is a member selected from the group consisting of:



1 6 (original): The method according to claim 1, wherein:
2 R_3 is a member selected from the group consisting of substituted alkyl and
3 substituted aryl groups.

1 7 (original): The method according to claim 6, wherein R_3 is a member selected
2 from the group consisting of:



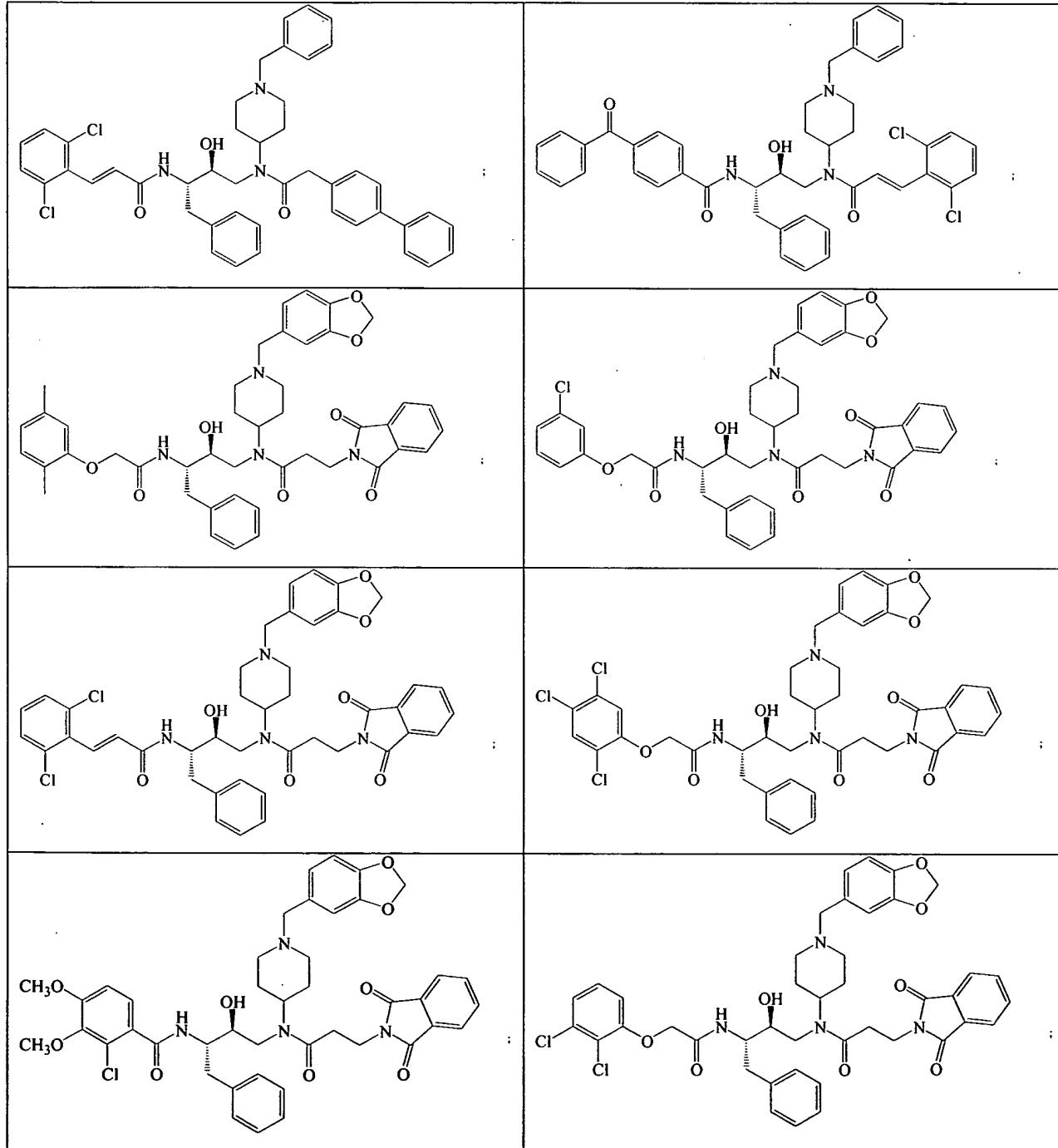
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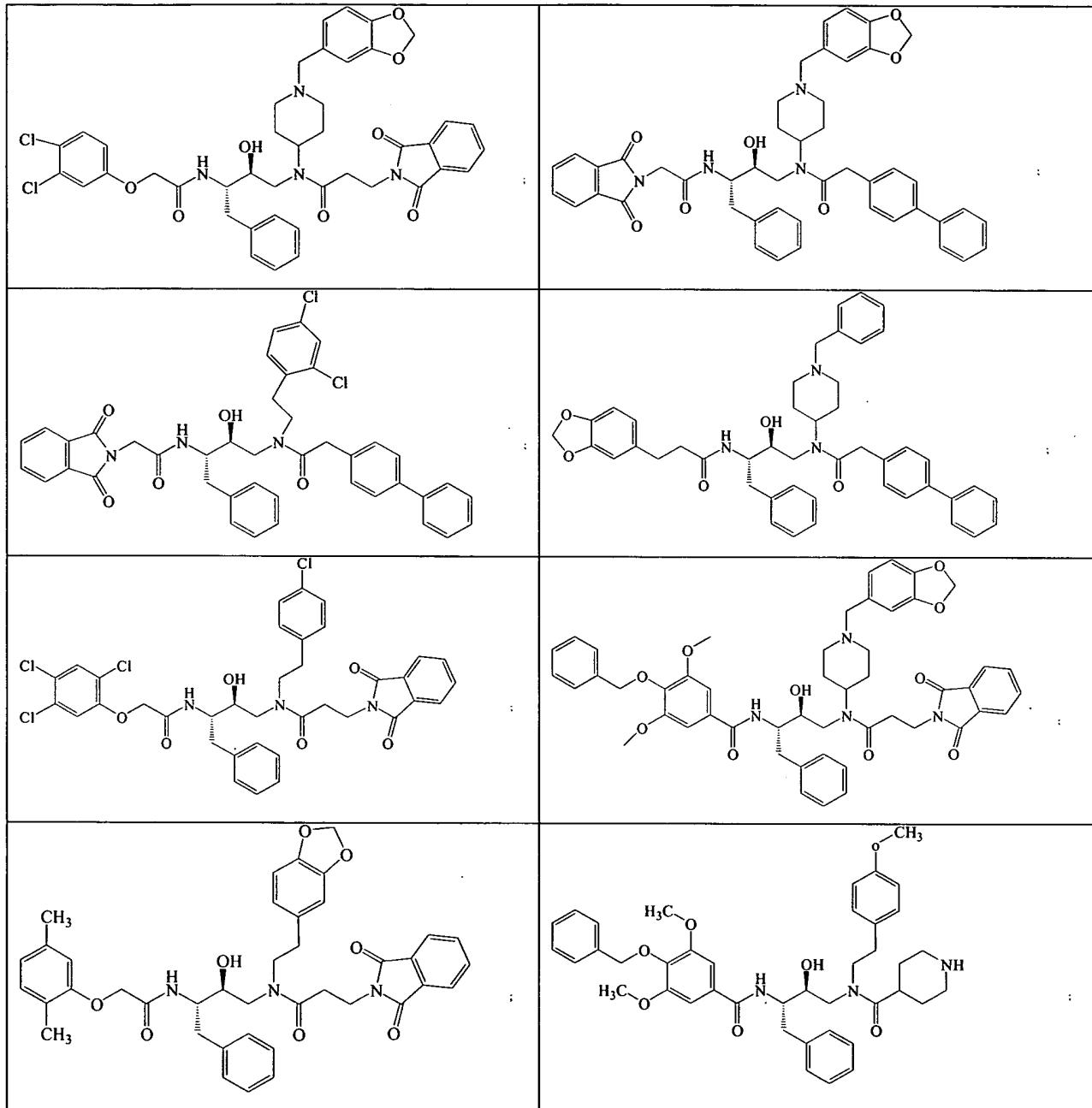
1 8 (original): The method according to claim 1, wherein R₅ and R₆ and the
2 carbons to which they are bound form an optionally substituted naphthalene ring.

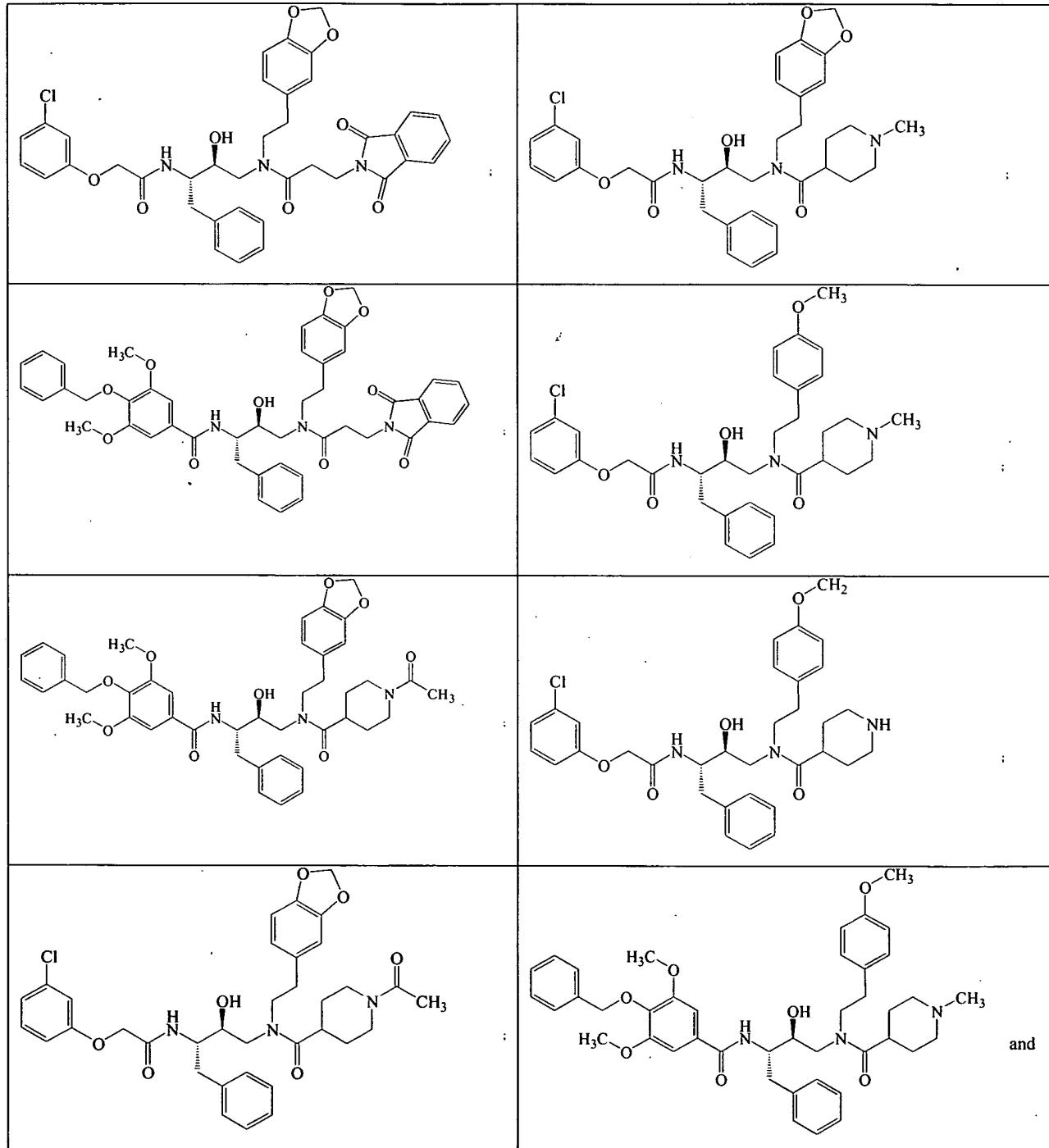
1 9 (original): The method according to claim 1, wherein R₅ and R₆ are both
2 hydrogen.

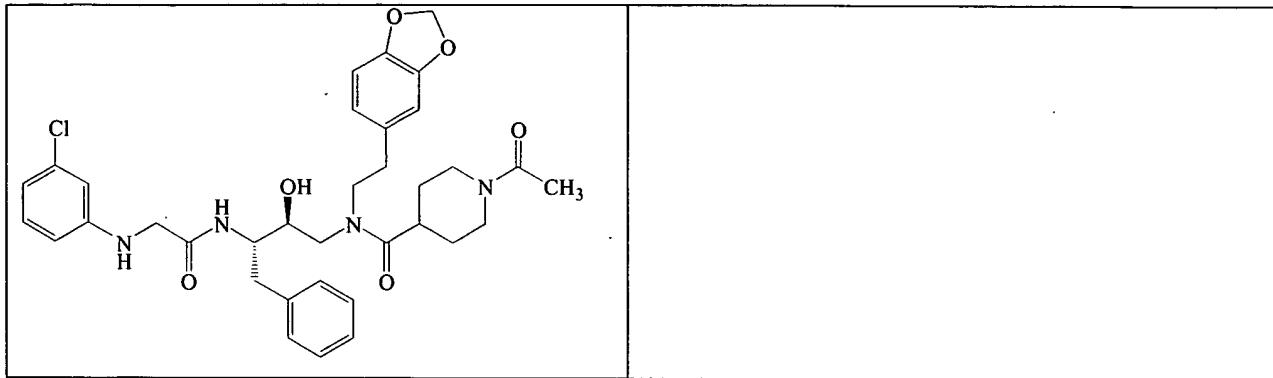
1 10 (original): The method in accordance with claim 1, wherein R₅ is hydrogen
2 and R₆ is meta or para to R₅ and is a member selected from the group consisting of halogen,
3 alkyl, substituted alkyl, aryl, substituted aryl, arylalkyl, substituted arylalkyl, aryloxyalkyl and
4 substituted aryloxyalkyl.

1 11 (original): The method according to claim 1, wherein said aspartyl protease
2 inhibitor is a member selected from the group consisting of:





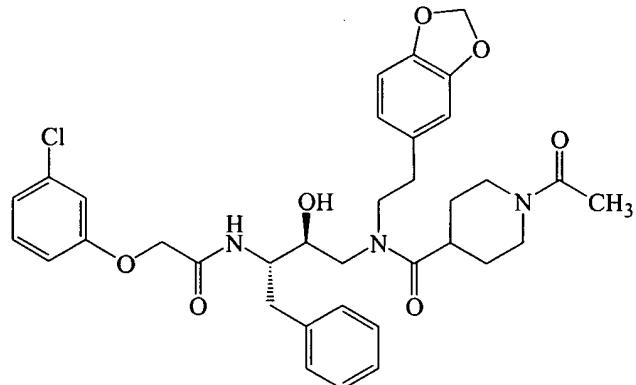




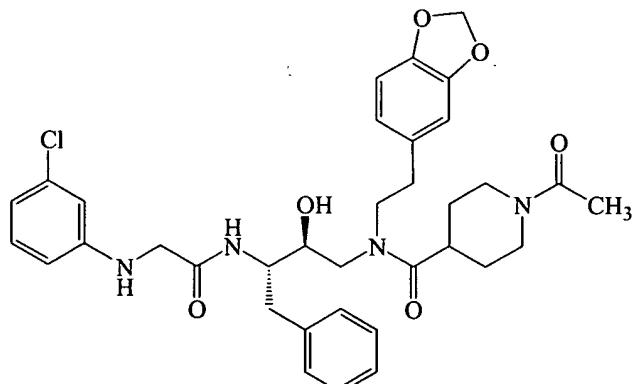
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1 12 (original): The method according to claim 1, wherein said aspartyl protease
2 inhibitor is a member selected from the group consisting of:



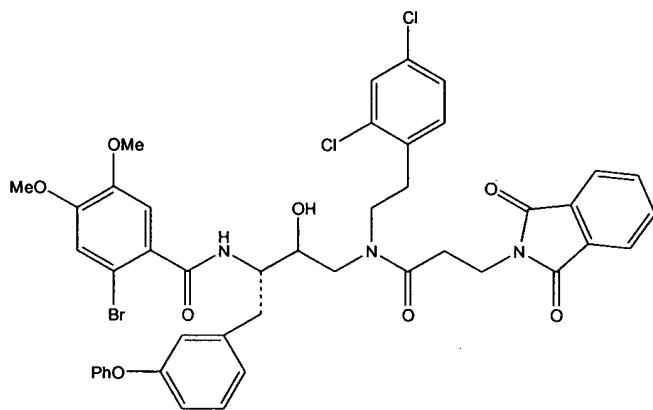
3 and



4

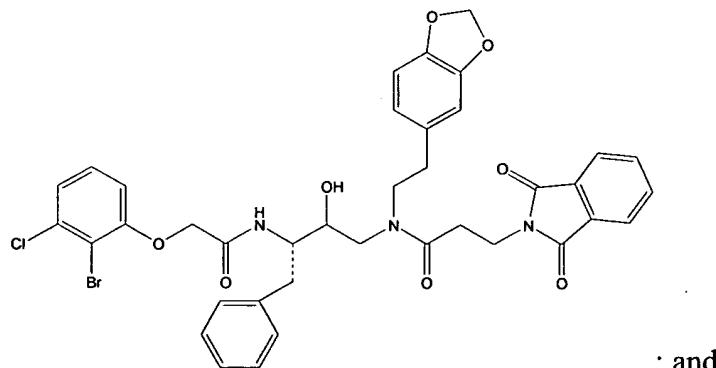
1 13 (currently amended): The method in accordance with claim 1, wherein said
2 aspartyl protease inhibitor is a member selected from the group consisting of ~~CEL5-A, CEL5-G~~
3 and ~~EA-1, which are illustrated in FIG. 12~~

4 CEL5-A having the following structure:



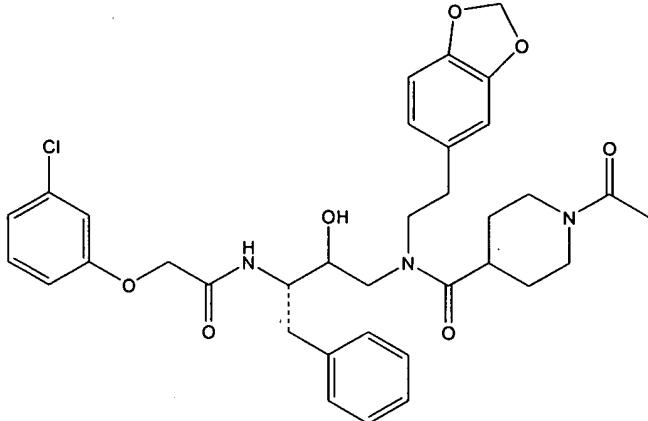
5 ;

6 CEL5G having the following structure:



7 ; and

8 EA 1 having the following structure:



9

1 14 (original): The method in accordance with claim 1, wherein said composition
2 is a body fluid.

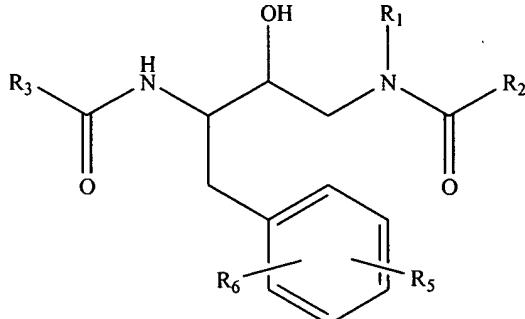
1 15 (currently amended): The method in accordance with claim [[13,]] 14,
2 wherein said body fluid is cerebral spinal fluid.

1 16 (original): The method in accordance with claim 1, whereby formation of
2 amyloidogenic A β peptides (A β) is decreased compared to the amount formed in the absence of
3 said aspartyl protease inhibitor.

1 17 (original): The method in accordance with claim 1, whereby formation of α -
2 sAPP is increased compared to the amount formed in the absence of said aspartyl protease
3 inhibitor.

1 18 (original): The method in accordance with claim 1, wherein the modulation is
2 effected by modulating the activity of cathepsin D.

1 19 (currently amended): A method for modulating the processing of a tau-
2 protein (τ -protein), said method comprising contacting a composition containing said τ -protein
3 with an aspartyl protease inhibitor having the **general** formula:



4 (I)

5 wherein:

6 R₁, R₂ and R₃ are members independently selected from the group consisting of
7 alkyl, substituted alkyl, aryl, substituted aryl, arylalkyl, substituted
8 arylalkyl, aryloxyalkyl, substituted aryloxyalkyl, heteroaryl, substituted
9 heteroaryl, heteroarylalkyl, substituted heteroarylalkyl, heterocycles,
10 substituted heterocycles, heterocyclicalkyl and substituted
11 heterocyclicalkyl; and

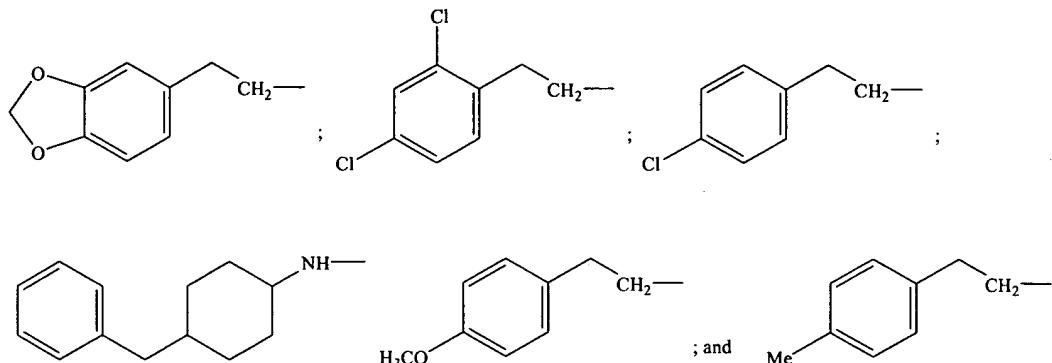
12 R₅ and R₆ are independently selected from the group consisting of hydrogen,
13 halogen, alkyl, substituted alkyl, aryl, substituted aryl, arylalkyl,
14 substituted arylalkyl, aryloxyalkyl and substituted aryloxyalkyl; or R₅ and
15 R₆ and the carbons to which they are bound join to form an optionally
16 substituted carbocyclic or heterocyclic fused ring system having a total of
17 9- or 10-ring atoms within said fused ring system.

1 20 (original): The method according to claim 19, wherein:

2 R₁ is a member selected from the group consisting of substituted alkylaryl,
3 substituted aryl, substituted alkyl and substituted heterocyclic groups.

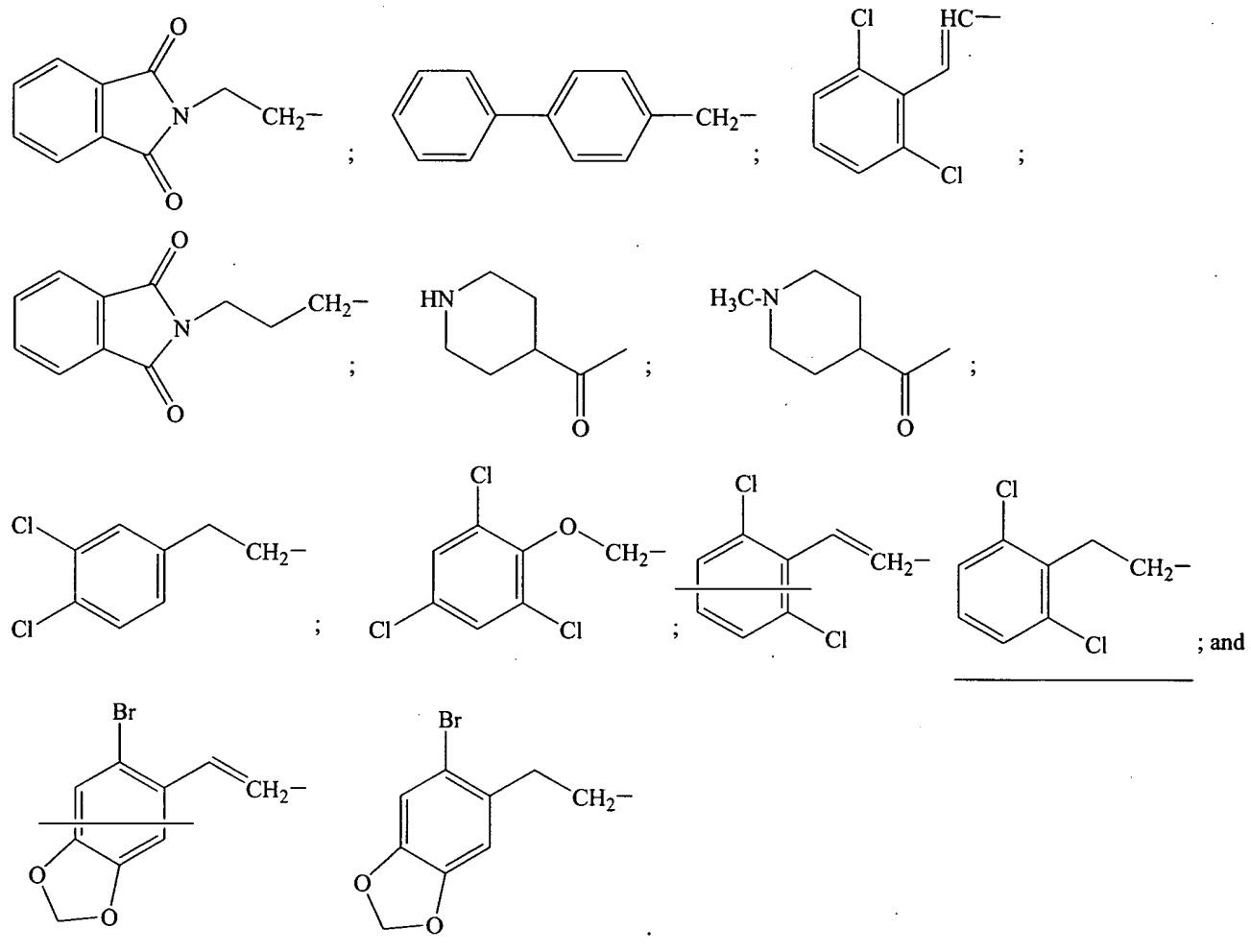
1 21 (original): The method according to claim 20, wherein:

2 R₁ is a member selected from the group consisting of:



1 22 (original): The method according to claim 19, wherein:
2 R₂ is a member selected from the group consisting of substituted alkyl,
3 heterocyclic and substituted heterocyclic groups.

1 23 (currently amended): The method according to claim 22, wherein R₂ is a
2 member selected from the group consisting of:



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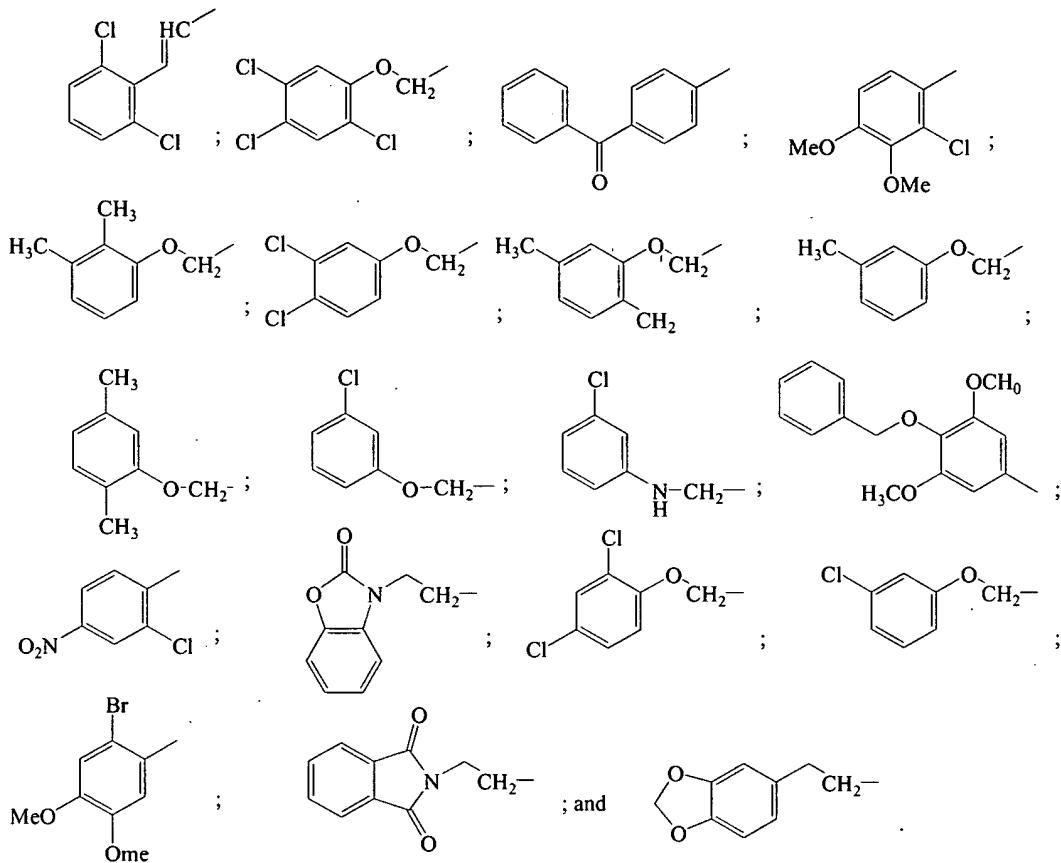
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24 (original): The method according to claim 19, wherein:

R₃ is a member selected from the group consisting of substituted alkyl and substituted aryl groups.

1 25 (original): The method according to claim 24, wherein R_3 is a member
2 selected from the group consisting of:

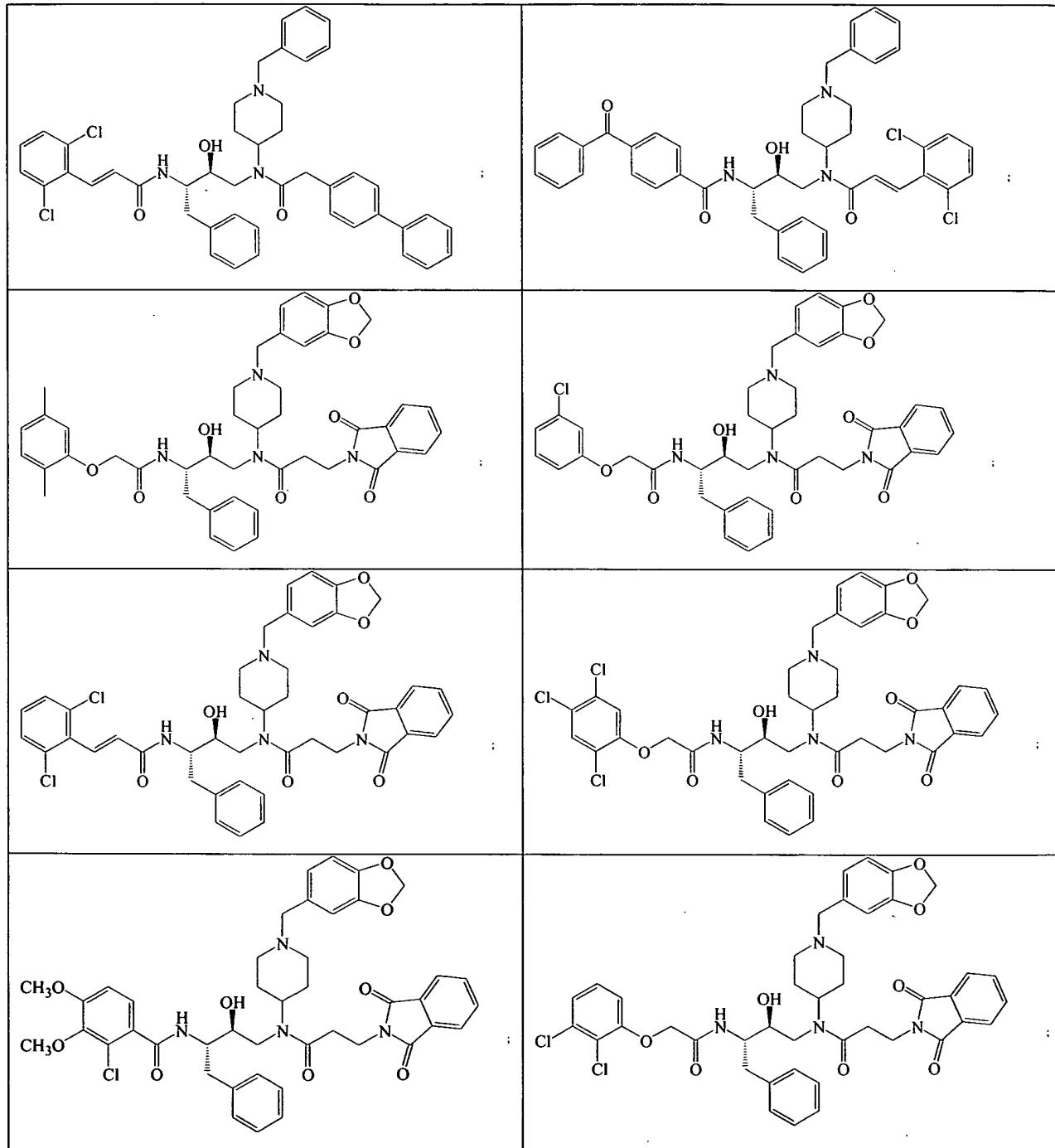


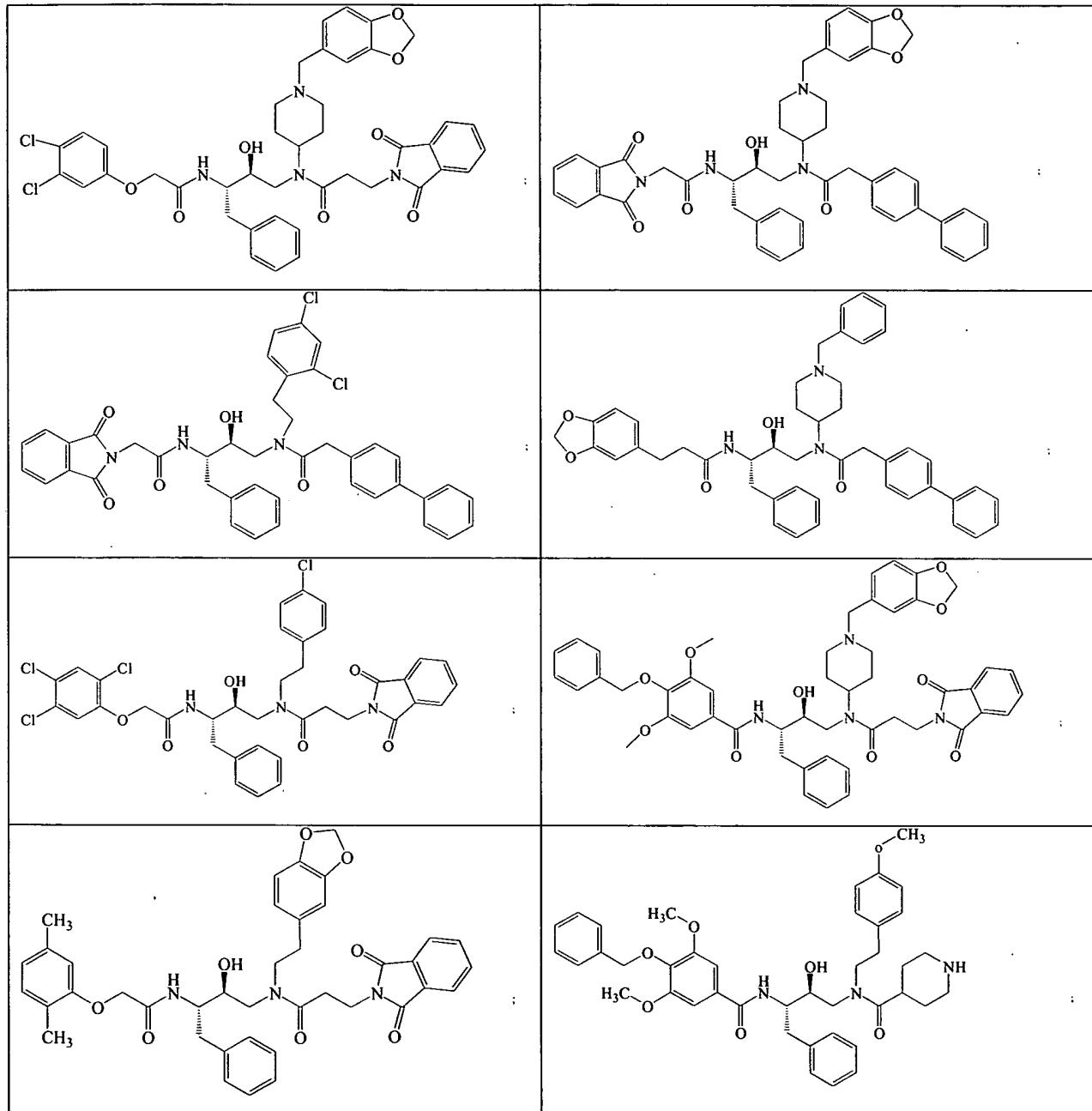
1 26 (original): The method according to claim 19, wherein R₅ and R₆ and the
2 carbons to which they are bound form an optionally substituted naphthalene ring.

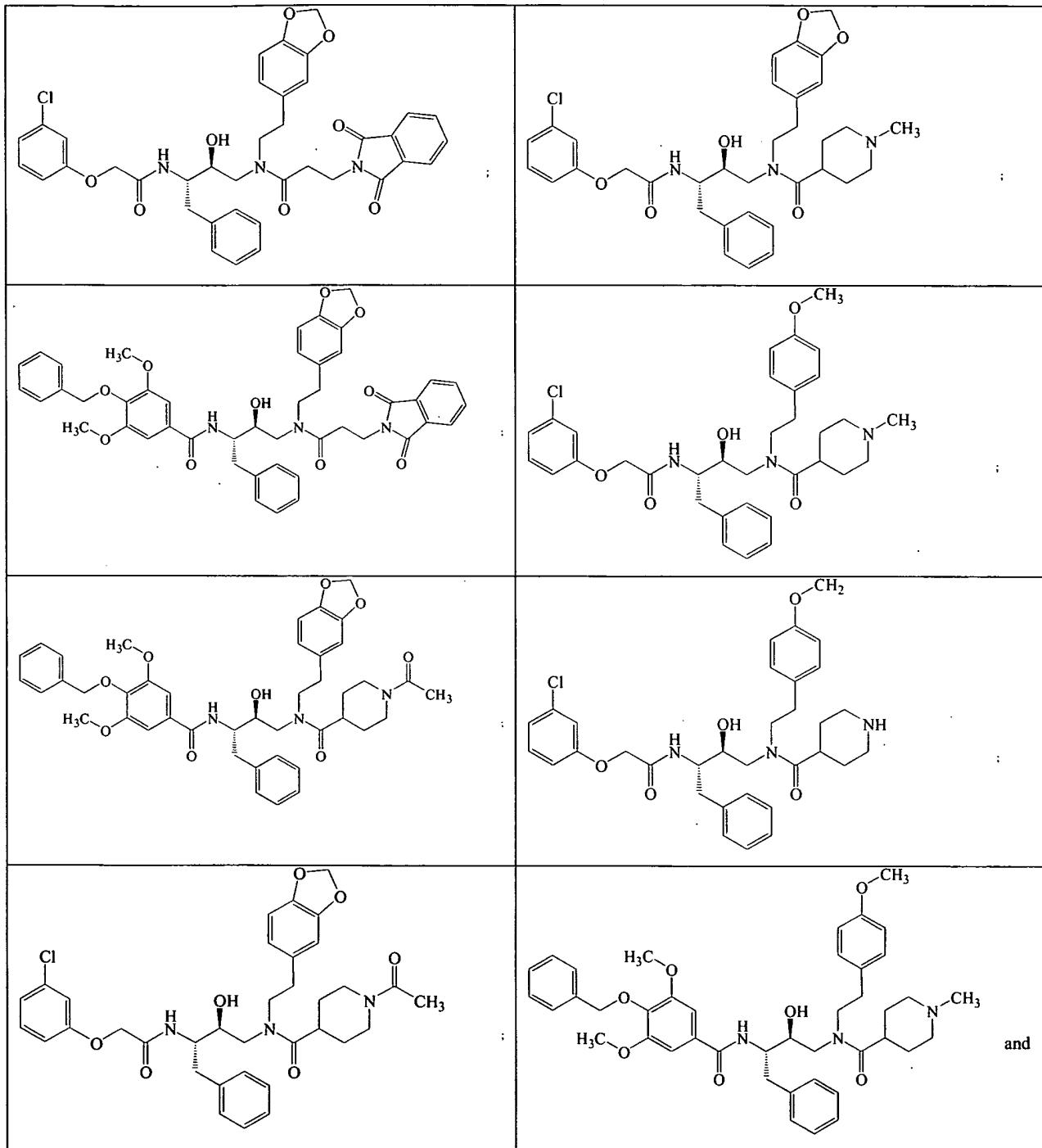
1 27 (original): The method according to claim 19, wherein R₅ and R₆ are both
2 hydrogen.

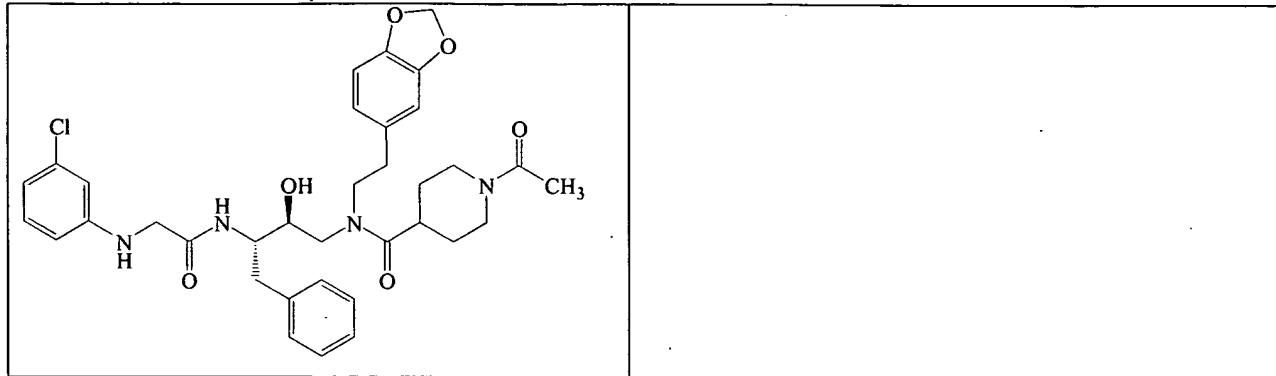
1 28 (original): The method in accordance with claim 19, wherein R₅ is hydrogen
2 and R₆ is meta or para to R₅ and is a member selected from the group consisting of halogen,
3 alkyl, substituted alkyl, aryl, substituted aryl, arylalkyl, substituted arylalkyl, aryloxyalkyl and
4 substituted aryloxyalkyl.

1 29 (original): The method according to claim 19, wherein said aspartyl protease
2 inhibitor is a member selected from the group consisting of:



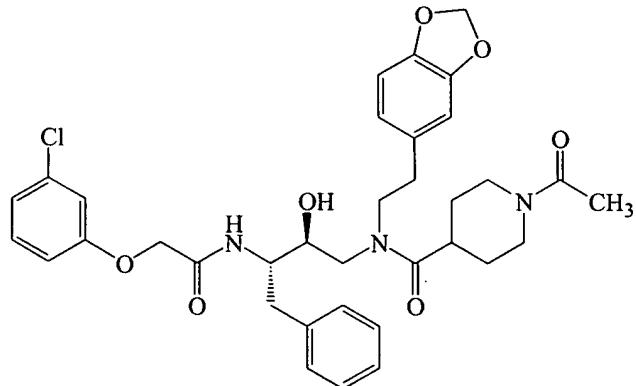




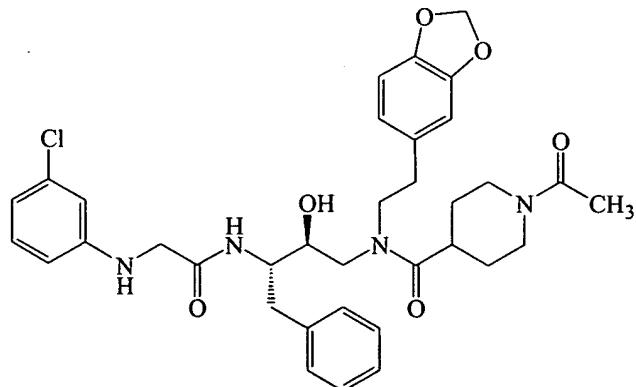


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1 30 (original): The method according to claim 19, wherein said aspartyl protease
2 inhibitor is a member selected from the group consisting of:



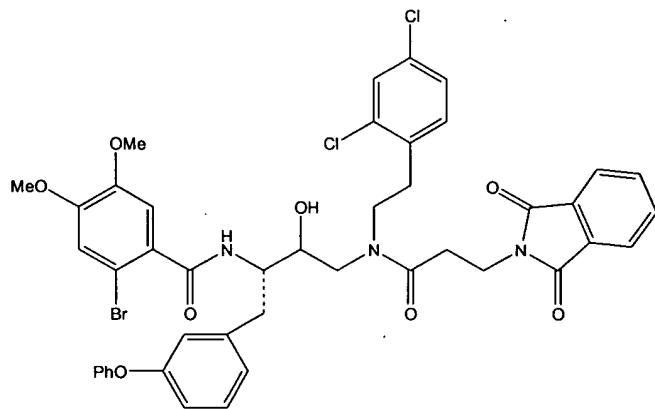
3 and



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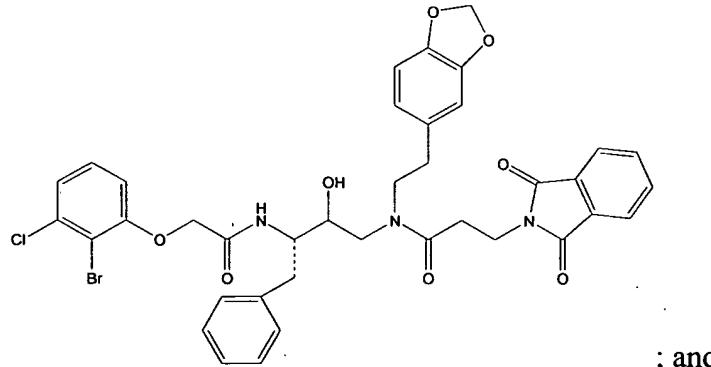
1 31 (currently amended): The method in accordance with claim 19, wherein said
2 aspartyl protease inhibitor is a member selected from the group consisting of ~~CEL5-A, CEL5-G~~
3 and ~~EA-1, which are illustrated in FIG. 12~~

4 CEL5-A having the following structure:



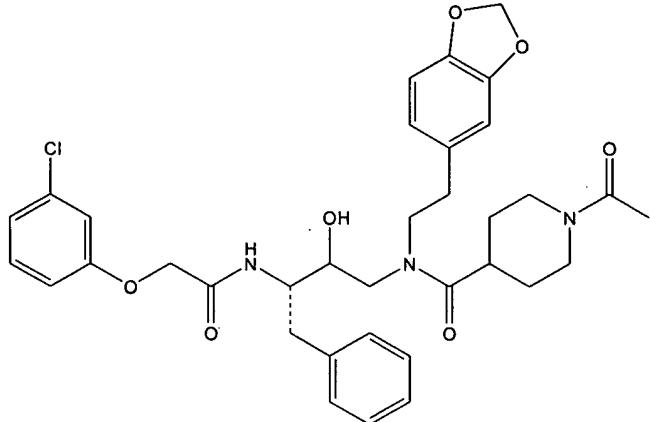
5 ;

6 CEL5G having the following structure:



7 ; and

8 EA 1 having the following structure:



9

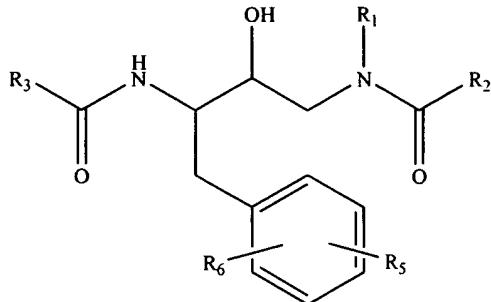
1 32 (original): The method in accordance with claim 19, wherein said
2 composition is a body fluid.

1 33 (currently amended): The method in accordance with claim [[31,]] 32,
2 wherein said body fluid is cerebral spinal fluid.

1 34 (original): The method in accordance with claim 19, whereby formation of τ -
2 fragments is decreased compared to the amount formed in the absence of said aspartyl protease
3 inhibitor.

1 35 (original): The method in accordance with claim 19, wherein the modulation
2 is effected by modulating the activity of cathepsin D.

1 36 (currently amended): A method for treating a neurodegenerative disorder,
2 said method comprising: administering to a mammal a therapeutically effective amount of an
3 aspartyl protease inhibitor having the **general** formula:



(I)

4 wherein:

5 R₁, R₂ and R₃ are members independently selected from the group consisting of
6 alkyl, substituted alkyl, aryl, substituted aryl, arylalkyl, substituted
7 arylalkyl, aryloxyalkyl, substituted aryloxyalkyl, heteroaryl, substituted
8 heteroaryl, heteroarylalkyl, substituted heteroarylalkyl, heterocycles,
9 substituted heterocycles, heterocyclicalkyl and substituted
10 heterocyclicalkyl; and

11 R₅ and R₆ are independently selected from the group consisting of hydrogen,
12 halogen, alkyl, substituted alkyl, aryl, substituted aryl, arylalkyl,
13 substituted arylalkyl, aryloxyalkyl and substituted aryloxyalkyl; or R₅ and
14 R₆ and the carbons to which they are bound join to form an optionally
15 substituted carbocyclic or heterocyclic fused ring system having a total of
16 9- or 10-ring atoms within said fused ring system; and
17 a pharmaceutically acceptable carrier,

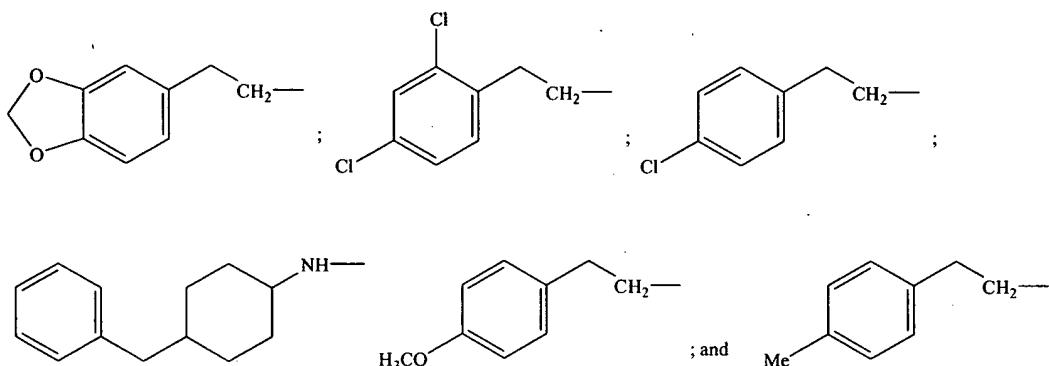
18 wherein: said neurodegenerative disorder is characterized by the accumulation of amyloid
19 plaques or by the accumulation of the accumulation of τ -fragments.

20
37-38 (canceled)

1 39 (original): The method in accordance with claim 36, wherein said
2 neurodegenerative disorder is a member selected from the group consisting of Alzheimer's
3 disease, Parkinson's disease, cognition defects, Downs Syndrome, cerebral hemorrhage with
4 amyloidosis, dementia and head trauma.

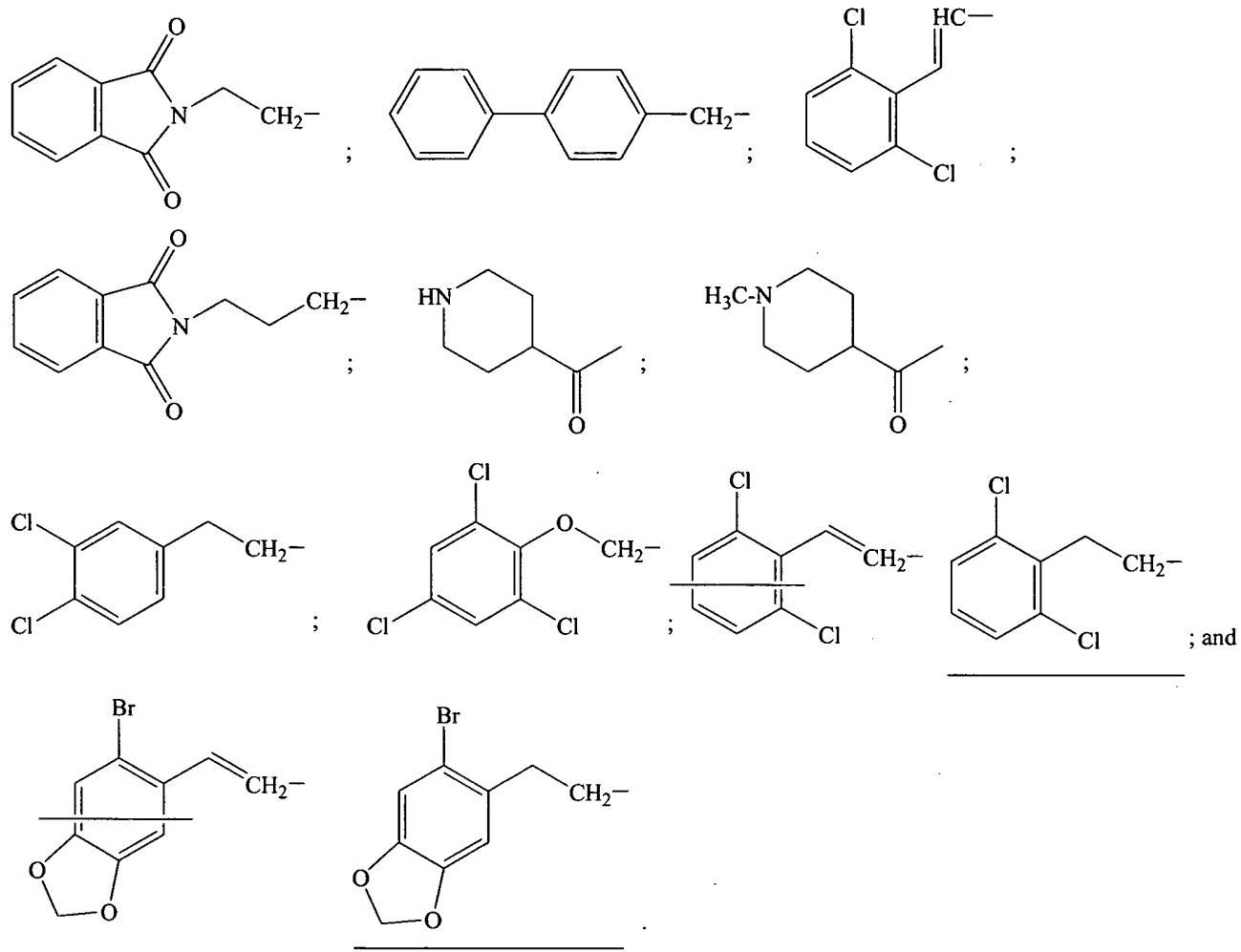
1 40 (original): The method according to claim 36, wherein:
2 R₁ is a member selected from the group consisting of substituted alkylaryl,
3 substituted aryl, substituted alkyl and substituted heterocyclic groups.

1 41 (original): The method according to claim 40, wherein:
2 R₁ is a member selected from the group consisting of:



1 42 (original): The method according to claim 36, wherein:
2 R₂ is a member selected from the group consisting of substituted alkyl,
3 heterocyclic and substituted heterocyclic groups.

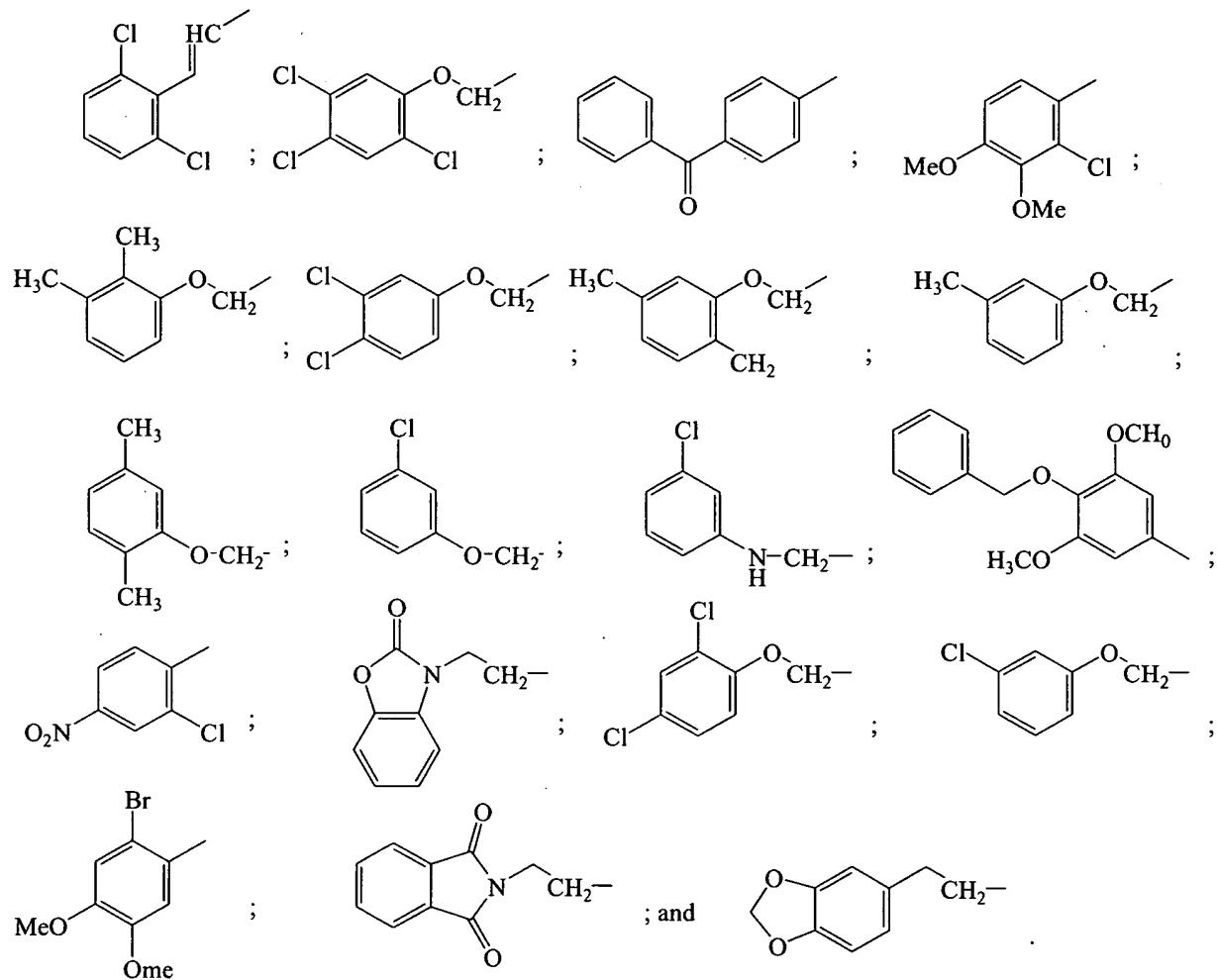
1 43 (currently amended): The method according to claim 42, wherein R₂ is a
2 member selected from the group consisting of:



1 44 (original): The method according to claim 36, wherein:

2 R₃ is a member selected from the group consisting of substituted alkyl and
3 substituted aryl groups.

1 45 (original): The method according to claim 44, wherein R₃ is a member
2 selected from the group consisting of:

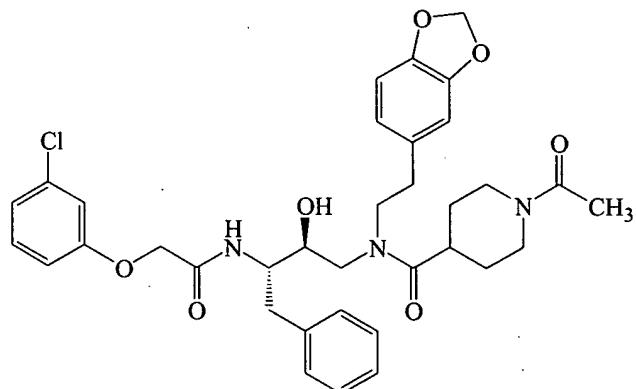


1 46 (original): The method according to claim 36, wherein R_5 and R_6 and the
2 carbons to which they are bound form an optionally substituted naphthalene ring.

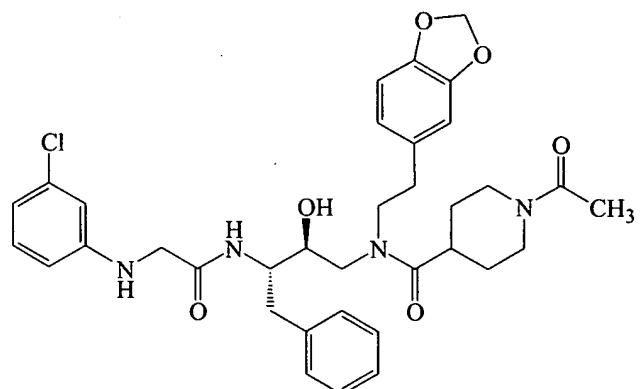
1 47 (original): The method according to claim 36, wherein R_5 and R_6 are both
2 hydrogen.

1 48 (original): The method in accordance with claim 36, wherein R₅ is hydrogen
2 and R₆ is meta or para to R₅ and is a member selected from the group consisting of halogen,
3 alkyl, substituted alkyl, aryl, substituted aryl, arylalkyl, substituted arylalkyl, aryloxyalkyl and
4 substituted aryloxyalkyl.

1 49 (original): The method in accordance with claim 36, wherein said aspartyl
2 protease inhibitor is a member selected from the group consisting of:



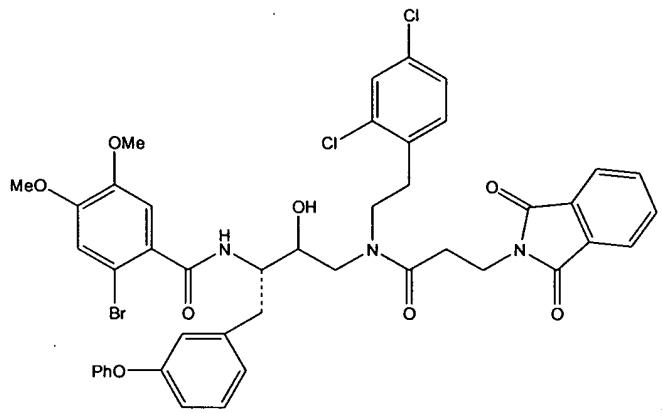
and



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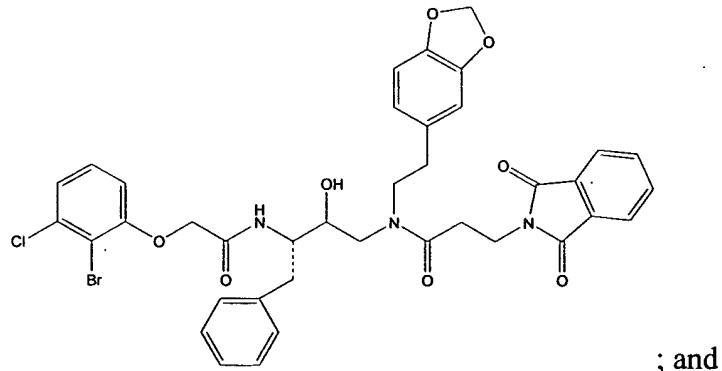
50 (currently amended): The method in accordance with claim 36, wherein said aspartyl protease inhibitor is a member selected from the group consisting of ~~CEL5-A, CEL5-G and EA-1, which are illustrated in FIG. 12~~

CEL5-A having the following structure:



;

CEL5G having the following structure:



; and

EA 1 having the following structure:

Appl. No. 10/774,262
Amdt. dated June 2, 2005
Reply to Office Action of December 2, 2004

PATENT

